Page 3

Amendments to the Claims:

- 1. (Currently amended) A method of preparing a chemically modified hemoglobin solution comprising an endogenous antioxidant enzyme, said method comprising:
- a) contacting a stroma free hemoglobin solution with at least one filtration means-filter, wherein a first filter filtration means retains viral particles and allows passage of a filtrate comprising a hemoglobin polypeptide and an endogenous antioxidant enzyme enzymes and the filtrate is substantially free of viral contamination;
 - b) chemically modifying the filtrate with an agent; and,
- c) isolating a composition comprising a the chemically modified hemoglobin solution and the endogenous antioxidant enzyme, wherein at least one of the endogenous antioxidant polypoptide enzymes retains enzymatic activity.
- 2. (Currently amended) The method of claim 1, wherein at least one of the endogenous antioxidant enzymes retaining enzymatic activity is selected from the group consisting of a superoxide dismutase, a catalase, and a glutathione peroxidase.
- 3. (Currently amended) The method of claim 1, wherein said first filter filtration means allows the passage of at least 50% of the endogenous antioxidant enzymes present in the stroma free hemoglobin solution.
- 4. (Currently amended) The method of claim 1, wherein the <u>first filter</u> filtration means comprises an AG-Technology a 500,000 molecular weight cutoff filter.
- 5. (Currently amended) The method of claim 1, wherein said first filter filtration-means reduces the passage of viral particles that are between about 200-25 nm in size.
- 6. (Currently amended) The method of claim 1, wherein said first filter filtration-means reduces the passage of viral particles that are 80-100 nm in size.

Page 4

- 7. (Currently amended) The method of claim 1, wherein said first filter filtration means reduces the passage of viral particles that are between about 80-50 nm in size.
- 8. (Currently amended) The method of claim 1, where said first <u>filter</u> filtration means reduces the passage of viral particles that are between about 50-25 nm in size.
- 9. (Currently amended) The method of claim 5, wherein said first <u>filter</u> filtration means reduces the passage of said viral particles by about 3 to about 10 log units.
- 10. (Currently amended) The method of claim 1, wherein said first filter filtration means produces a filtrate having a viral load reduction of at least 3 log units.
- 11. (Currently amended) The method of claim 1 further comprising contacting the filtrate with at least-one a second filter filtration-means wherein said second filter filtration means allows the passage of the hemoglobin polypeptide and the endogenous antioxidant enzymes enzyme and retains virus particles.
 - 12. (Canceled)
- 13. (Currently amended) The method of claim 1, wherein the modifying agent is a bifunctional modifying agent.
- 14. (Currently amended) The method of claim 13, wherein said modifying agent is selected from the group consisting of <u>a</u> sebacyl chloride, <u>a</u> glutaraldehyde, <u>a</u> diasprin derivatives, <u>a</u> polyaldehydes, <u>a</u> polyoxyetheylene, <u>a</u> dextrans, and <u>an</u> insulin.
- 15. (Currently amended) The method of claim 13, wherein the modifying agent is a bifunctional polyoxyetheylene.

Page 5

- 16. (Currently amended) The method of claim 1, wherein the modifying agent is a mixture of a bifunctional and a monofunctional polyoxyethylene.
- 17. (Currently amended) The method of claim 15, wherein the <u>chemically</u> modified hemoglobin solution <u>comprising an endogenous antioxidant enzyme</u> is PHP.
- 18. (Currently amended) The method of claim 1, wherein the chemical modification chemically modifying said filtrate with an agent further comprises deoxygenation and pyridoxalation of the hemoglobia.
- 19. (Currently amended) The method of claim 1, wherein the viral contamination of said isolated chemically modified hemoglobin solution comprising an endogenous antioxidant enzyme comprises a hepatitis A viral titer of less than about 1 TCID₅₀ unit/ml.
- 20. (Original) The method of claim 1, wherein the chemically modified hemoglobin solution comprises about a 50% to about a 200% increase in endogenous red blood cell antioxidant activity per unit of hemoglobin found in red blood cells.
- 21. (Currently amended) A method of preparing a chemically modified hemoglobin solution comprising an endogenous antioxidant enzyme, said method consisting of:
- a) contacting a stroma free hemoglobin solution with at least one <u>filter</u> filtration means, wherein a first <u>filter filtration means</u> retains viral particles and allows passage of a filtrate comprising <u>a hemoglobin polypeptide</u> and <u>the</u> endogenous antioxidant <u>enzyme</u> enzymes and the filtrate is substantially free of viral contamination;
 - b) chemically modifying the filtrate with an agent; and,
- c) isolating a composition comprising the chemically modified hemoglobin solution and the endogenous antioxidant enzyme enzymes.
- 22. (Currently amended) A hemoglobin solution comprising a chemically modified hemoglobin solution comprising and at least one endogenous antioxidant enzyme, wherein said

Page 6

chemically modified hemoglohin solution modification comprises attachment of a POE linkage, said endogenous antioxidants antioxidant enzyme retains retain enzymatic activity, and said chemically modified hemoglobin solution is substantially free of a viral contamination.

- 23. (Currently amended) The <u>chemically</u> modified hemoglobin solution of claim 22, wherein said <u>chemically</u> modified hemoglobin <u>solution</u> is PHP.
- 24. (Currently amended) The chemically modified hemoglobin solution of claim 22, wherein the viral contamination of a viral particle of less than 70 nm in size is of said solution comprises a viral titer of less than about 1 TCID₅₀ unit/ml.
- 25. (Currently amended) The chemically modified hemoglobin solution of claim 24, wherein the viral contamination titer of said particles that are a viral particle that is 25-30 nm in size is less than about 1 TCID₅₀ unit/ml.
- 26. (Currently amended) The chemically modified hemoglobin solution of claim 25, wherein said viral particle is hepatitis A.
 - 27. (Canceled)
- 28. (Currently amended) The chemically modified hemoglobin solution of claim 27, wherein the viral particle is hepatitis A or hepatitis C.
- 29. (Currently amended) The chemically modified hemoglobin solution of claim 22, wherein said endogenous antioxidant enzyme is selected from the group consisting of a superoxide dismutase, a catalase, a hemoglobin peroxidase, and a glutathione peroxidase.
- 30. (Currently amended) The chemically modified hemoglobin solution of claim 22, wherein said solution contains comprises between a 50% to a 200% increase in antioxidant activity per unit of hemoglobin found in red blood cells.

Page 7

31. (Canceled)

- 32. (Currently amended) A method of decreasing the level of nitric oxide present in the circulation of a mammal, said method comprising, administering to a mammal in a need thereof a therapeutically effective amount of the <u>chemically</u> modified hemoglobin solution of claim 22 in a pharmaceutically acceptable carrier.
- 33. (Currently amended) The method of claim 32, wherein said <u>chemically</u> modified hemoglobin <u>solution</u> is administered to a mammal having systemic hypotension.
- 34. (Currently amended) The method of claim 32, wherein said chemically modified hemoglobin solution is administered to a mammal having septic shock.
- 35. (Currently amended) A method of treating red blood cell loss, said treatment comprising administering to a mammal in need thereof a therapeutically effective amount of the chemically modified hemoglobin solution of claim 22.